



Substitute for form 1449A/PTO

## INFORMATION DISCLOSURE STATEMENT BY APPLICANT

(Use as many sheets as necessary)

Sheet

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of 10

<i>Complete if Known</i>	
Application Number	10/698,099
Filing Date	October 31, 2003
First Named Inventor	Schenk, Dale B.
Art Unit	1648
Examiner Name	Michelle S. Horning
Attorney Docket Number	015270-008930US

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. <sup>1</sup>	Document Number Number Kind Code <sup>2</sup> ( <i>if known</i> )	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
/MH	1	US-7,138,255	11-21-2006	Vodyanoy et al.	
	2	US-2006/0259986	11-16-2006	Chilcote et al.	
	3	US-2006/0058233	03-16-2006	Schenk et al.	
	6	US-2005/0196818	09-08-2005	Chilcote et al.	
	7	US-2005/0198694	09-08-2005	Chilcote et al.	
	8	US-2005/0176078	08-11-2005	Allsop et al.	
	13	US-2005/0037013	02-17-2005	Schenk et al.	
	19	US-6,780,971	08-24-2004	Wolozin et al.	
	20	US-2004/0146521	07-29-2004	Schenk et al.	
	21	US-2004/0137523	07-15-2004	Vodyanoy et al.	
	22	US-2004/0136993	07-15-2004	Schenk et al.	
	25	US-6,710,226	03-23-2004	Schenk	
	32	US-2002/0160394 A1	10-31-2002	Wu	
	37	US-5,589,154	12-31-1996	Anderson	
	38	US-5,576,184	11-19-1996	Better et al.	
	39	US-4,883,666	11-28-1989	Sabel et al	

  

U.S. PATENT APPLICATIONS					
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	40	US-11/710,248	02-23-2007	Schenk et al.	
	41	US-11/697,646	04-06-2007	Schenk et al.	
	42	US-11/660,015	02-09-2007	Schenk et al.	
	43	US-10/850,570	05-19-2004	Chilcote et al.	
	44	US-60/518,140	11-08-2003	Chilcote et al.	
	45	US-60/471,929	05-19-2003	Chilcote et al.	

FOREIGN PATENT DOCUMENTS					
Examiner Initials*	Cite No. <sup>1</sup>	Foreign Patent Document Country Code <sup>3</sup> Number <sup>4</sup> Kind Code <sup>5</sup> ( <i>if known</i> )	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
	46	EP 0 613 007 A2	08-31-1994		<input type="checkbox"/>
	47	WO 07/021255 A1	02-22-2007	Chilcote et al.	<input type="checkbox"/>
	48	WO 07/012061 A2	01-25-2007	Schenk et al.	<input type="checkbox"/>
	49	WO 06/045037 A2	04-27-2006	Chilcote et al.	<input type="checkbox"/>

Examiner Signature	/Michelle Horning/	Date Considered	01/03/2008
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		Country Code <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)				
/MH/	50	WO	06/045037	A3	04-27-2006	Chilcote et al.		<input type="checkbox"/>
	51	WO	06/020581	A2 corrected version	02-23-2006	Schenk et al.		<input type="checkbox"/>
	52	WO	06/020581	A3	02-23-2006	Schenk et al.		<input type="checkbox"/>
	55	WO	05/047860	A3	05-26-2005	Chilcote et al..		<input type="checkbox"/>
	57	WO	04/041067	A3	05-21-2004	Schenk et al.		<input type="checkbox"/>
	59	WO	04/009625	A3	01-29-2004	El-Agnaf et al.		<input type="checkbox"/>
	184	WO	01/053457	A3	07-26-2001	University of Connecticut Health Center		<input type="checkbox"/>
	186	WO	01/06989	A3	02-01-2001	Abgenix, Inc.		<input type="checkbox"/>
	61	WO	00/72880	A3	12-07-2000	Schenk et al.		<input type="checkbox"/>
	63	WO	00/072876	A3	12-07-2000	Schenk		<input type="checkbox"/>
	64	WO	00/18917	A2 corrected version	04-06-2000	Amgen Inc.		<input type="checkbox"/>
	65	WO	00/18917	A3 corrected version	04-06-2000	Amgen Inc.		<input type="checkbox"/>
	67	WO	99/050300	A1	10-07-1999	Trojanowski et al.		<input type="checkbox"/>
	71	WO	99/006545	A2 corrected version	02-11-1999	Max Plank Institute		<input type="checkbox"/>
	72	WO	99/006545	A3 corrected version	02-11-1999	Max Plank Institute		<input type="checkbox"/>
	73	WO	95/006407	A1	03-09-1995	Masliah		<input type="checkbox"/>



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				<i>Examiner Name</i>	Michelle S. Horning
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/MH/	75	ABBAS et al., <u>Cellular and Molecular Immunology</u> , 522-523 (Elsevier Saunders) (5th Ed. Updated Ed., 2005).			<input type="checkbox"/>
	76	BALES et al., "Cholinergic dysfunction in a mouse model of Alzheimer disease is reversed by an anti-A $\beta$ antibody," <u>J. Clin. Invest.</u> , 116(3):825-832 (2006).			<input type="checkbox"/>
	187	BARD et al., "Peripherally administered antibodies against amyloid $\beta$ -peptide enter the central nervous system and reduce pathology in a mouse model of Alzheimer disease," <u>Nature Medicine</u> , 6(8):916-919 (2000).			<input type="checkbox"/>
	77	BENNETT et al., "Degradation of $\alpha$ -Synuclein by Proteasome," <u>J. Biol. Chem.</u> , 274(48):33855-33858 (1999).			<input type="checkbox"/>
	78	BROOKS et al., "Synuclein proteins and Alzheimer's disease," <u>Trends Neurosci.</u> , 17(10):404-405 (1994).			<input type="checkbox"/>
	80	CHEN et al., "Neurodegenerative Alzheimer-like pathology in PDAPP 717V $\rightarrow$ F transgenic mice," <u>Progress in Brain Research</u> , 117:327-337 (1998).			<input type="checkbox"/>
	81	CLAYTON et al., "Synucleins in Synaptic Plasticity and Neurodegenerative Disorders," <u>J. Neurosci. Res.</u> , 58:120-129 (1999).			<input type="checkbox"/>
	82	CLAYTON et al., "The synucleins: a family of proteins involved in synaptic function, plasticity, neurodegeneration, and disease," <u>Trends Neurosci.</u> , 21(6):249-254 (1998).			<input type="checkbox"/>
	83	CLELAND et al., "Isomerization and Formulation Stability of the Vaccine Adjuvant QS-21," <u>J. of Pharm Sci.</u> , 85(1): 22-28 (1996).			<input type="checkbox"/>
	85	CROWTHER et al., "Synthetic filaments assembled from C-terminally truncated $\alpha$ -synuclein," <u>FEBS Letters</u> , 436:309-312 (1998).			<input type="checkbox"/>

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				<i>Examiner Name</i>	Michelle S. Horning
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/MH/	88	Dictionary.com definition of "prophylactic", pages 1-3 downloaded from internet 10/12/05.	<input type="checkbox"/>
	90	DI MONTE et al., "Environmental Factors in Parkinson's Disease," <u>Neurotoxicology</u> , 23: 487-502 (2002).	<input type="checkbox"/>
	191	DIXON, C. et al., "Alpha-Synuclein Targets the Plasma Membrane via the Secretory Pathway and Induces Toxicity in Yeast," <u>Genetics</u> , 2005 May;170(1):47-59. Epub 2005 Mar 2.	<input type="checkbox"/>
	178	EL-AGNAF et al., " $\alpha$ -Synuclein implicated in Parkinson's disease is present in extracellular biological fluids, including human plasma," <u>FASEB J.</u> , 17(3):1945-1947 (2003).	<input type="checkbox"/>
	91	EL-AGNAF et al., " $\alpha$ -Synuclein implicated in Parkinson's disease is present in extracellular biological fluids, including human plasma," <u>FASEB J. express article 10.1096/fj.03-0098fie</u> , Published online August 15, 2003.	<input type="checkbox"/>
	94	ELIEZER, D. et al., "Conformational Properties of Alpha-Synuclein in its Free and Lipid-associated States," <u>Journal of Molecular Biology</u> , 307(4):1061-1073 (2001).	<input type="checkbox"/>
	95	ELLIS et al., " $\alpha$ -Synuclein is Phosphorylated by Members of the Src Family of Protein-tyrosine Kinases," <u>J. Biol. Chem.</u> , 276(6):3879-3884 (2001).	<input type="checkbox"/>
	96	EMADI, S. et al., "Inhibiting Aggregation of Alpha-Synuclein with Human Single Chain Antibody Fragments," <u>Biochemistry</u> , 43(10):2871-2878 (2004).	<input type="checkbox"/>
	98	EP 04776059.0 European Supplementary Search Report completed 06/13/2006.	<input type="checkbox"/>
	192	GAMES et al., "Prevention and Reduction of AD-type Pathology in PDAPP Mice Immunized with A $\beta$ 1-42," <u>Annals of the New York Academy of Science</u> , 920:274-284 (2000).	<input type="checkbox"/>

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/MH/	193	GARZON, J. et al., "Transport of CSF antibodies to G-Alpha subunits across neural membranes requires binding to the target protein and protein kinase C activity," <u>Molecular Brain Research</u> , 65(2):151-166 (1999).			<input type="checkbox"/>
	106	GIASSON et al., "Mutant and Wild Type Human $\alpha$ -Synucleins Assemble into Elongated Filaments with Distinct Morphologies <i>in Vitro</i> ," <u>J. Biol. Chem.</u> , 274(12):7619-7622 (1999).			<input type="checkbox"/>
	108	GOLDSTEINS et al., "Exposure of cryptic epitopes on transthyretin only in amyloid and in amyloidogenic mutants," <u>PNAS</u> , 96:3108-3113 (1999).			<input type="checkbox"/>
	110	HAMBURGER, A.W. et al., "Isolation and characterization of monoclonal antibodies reactive with endothelial cells," <u>Tissue &amp; Cell</u> , 17(4): 451-459 (1985).			<input type="checkbox"/>
	112	HARTMAN et al., "Treatment with an Amyloid- $\beta$ Antibody Ameliorates Plaque Load, Learning Deficits, and Hippocampal Long-Term Potentiation in a Mouse Model of Alzheimer's Disease," <u>Journal of Neuroscience</u> , 25:6213-6220 (2005).			<input type="checkbox"/>
	113	HOOPER et al., <u>Cellular Peptidases in Immune Functions and Diseases 2</u> , (Langer and Ansorge, Eds., Plenum Publishers) 379-390 (2000).			<input type="checkbox"/>
	114	HOYER, W. et al., "Dependence of alpha-Synuclein Aggregate Morphology on Solution Conditions," <u>J. Mol. Biol.</u> , 322:383-393 (2002).			<input type="checkbox"/>
	179	IWAI, "Properties of NACP/alpha-synuclein and its role in Alzheimer's disease," <u>Molecular Basis of Disease</u> , 1502(1): 95-109 (2000).			<input type="checkbox"/>
	116	IWATSUBO, T. et al., "Purification and Characterization of Lewy Bodies from the Brains of Patients with Diffuse Lewy Body Disease," <u>Am J Pathol.</u> , 148(5):1517-1529 (1996).			<input type="checkbox"/>
	117	JAKES et al., "Epitope mapping of LB509, a monoclonal antibody directed against human $\alpha$ -synuclein," <u>Neurosci. Letts.</u> , 269:13-16 (1999).			<input type="checkbox"/>

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/MH/	120	KIM, T. D. et al., "Structural and Functional Implications of C-Terminal Regions of $\alpha$ -Synuclein," <u>Biochemistry</u> , 41:13782-13790 (2002).			<input type="checkbox"/>
	121	KIM, T.D. et al., "Structural Changes in $\alpha$ -Synuclein Affect its Chaperone-like Activity in Vitro," <u>Protein Science</u> , 9:2489-2496 (2000).			<input type="checkbox"/>
	123	KUBY J., eds., <u>Immunology</u> , pp. 92-97 and 131 (W.H. Freeman & Co., New York) (3rd Edition, 1997).			<input type="checkbox"/>
	124	KUBY J., eds., <u>Immunology</u> , pp. 156-158 (W.H. Freeman & Co., New York) (3rd Edition, 1997).			<input type="checkbox"/>
	199	LANSBURY JR., P. T., "Evolution of amyloid: What normal protein folding may tell us about fibrillogenesis and disease," <u>Proc Natl Acad Sci</u> , 96(7): 3342-3344 (1999).			<input type="checkbox"/>
	125	LEE et al., "Formation and Removal of $\alpha$ -Synuclein Aggregates in Cells Exposed to Mitochondrial Inhibitors," <u>J. Biol. Chem.</u> , 277(7):5411-5417 (2002).			<input type="checkbox"/>
	126	LEE et al., "Human $\alpha$ -synuclein-harboring familial Parkinson's disease-linked Ala-53 → Thr mutation causes neurodegenerative disease with $\alpha$ -synuclein aggregation in transgenic mice," <u>PNAS</u> , 99:8968-8973 (2002).			<input type="checkbox"/>
	200	LEMERE, C. A. et al., Amyloid-Beta Immunization in Alzheimer's Disease Transgenic Mouse Models and Wildtype Mice," <u>Neurochem Res.</u> , 28(7):1017-27.2003).			<input type="checkbox"/>
	201	LUTHI-CARTER, R., "Progress towards a Vaccine for Huntington's Disease," <u>Mol Ther.</u> , 7(5, Pt 1):569-70 (2003).			<input type="checkbox"/>
	203	MCLEAN, et al., "Membrane Association and Protein Conformation of Alpha-Synuclein in Intact Neurons," <u>J Biol Chem.</u> , 275(12):8812-6 (2000).			<input type="checkbox"/>
	133	Merriam-Webster online medical dictionary, entry for "cure", accessed September 5, 2006.			<input type="checkbox"/>

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/MH/	134	MISHIZEN-EBERZ et al., "Distinct cleavage patterns of normal and pathologic forms of $\alpha$ -synuclein by calpain I <i>in vitro</i> ," <u>J. Neurochemistry</u> , 86:836-847 (2003).			<input type="checkbox"/>
	204	MORGAN, et al., " $\text{A}\beta$ peptide vaccination prevents memory loss in an animal model of Alzheimer's disease," <u>Nature</u> , 408(6815):982-5 (2000).			<input type="checkbox"/>
	136	OKOCHI, M. "Constitutive Phosphorylation of the Parkinson's Disease Associated $\alpha$ -Synuclein," <u>J. Biol. Chem.</u> , 275(1): 390-397 (2000).			<input type="checkbox"/>
	137	PALHA et al., "Antibody recognition of amyloidogenic transthyretin variants in serum of patients with familial amyloidotic polyneuropathy," <u>J. Mol. Med.</u> , 78:703-707 (2001).			<input type="checkbox"/>
	138	PCT/US05/37875 International Preliminary Report on Patentability Chapter 1 issued 04/24/2007 with Written Opinion			<input type="checkbox"/>
	139	PCT/US05/28166 International Preliminary Report on Patentability Chapter 1 issued 02/13/2007 with Written Opinion			<input type="checkbox"/>
	140	PCT/US04/37444 International Preliminary Report on Patentability Chapter 1 issued 06/19/2007 with Written Opinion			<input type="checkbox"/>
	141	PCT/US04/015836 International Preliminary Report on Patentability Chapter 1 issued 11/25/2005 with Written Opinion			<input type="checkbox"/>
	142	PCT/US00/015239 International Preliminary Examination Report dated 08/13/2001			<input type="checkbox"/>
	145	QUE et al., "Effect of Carrier Selection on Immunogenicity of Protein Conjugate Vaccines against <i>Plasmodium falciparum</i> Circumsporozoites," <u>Infection and Immunity</u> , 56(10): 2645-2649 (1988).			<input type="checkbox"/>
↓	147	SCHENK, D., "Amyloid- $\beta$ immunotherapy for Alzheimer's disease: the end of the beginning," <u>Nature Reviews</u> , 3:824-828 (2002).			<input type="checkbox"/>
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/MH/	149	SIGURDSSON et al., "Immunization Delays the Onset of Prion Disease in Mice," <u>American Journal of Pathology</u> , 161:13-17 (2002).			<input type="checkbox"/>
	150	SIGURDSSON et al., "Anti-prion antibodies for prophylaxis following prion exposure in mice," <u>Neurosciences Letters</u> , 336:185-187 (2003).			<input type="checkbox"/>
	151	SIPE, "Amyloidosis," <u>Annu. Rev. Biochem.</u> , 61:947-975 (1992).			<input type="checkbox"/>
	152	SKIPPER et al., "Parkinson's Genetics: molecular Insights for the New Millennium," <u>Neurotoxicology</u> , 23: 503-514 (2002).			<input type="checkbox"/>
	155	SMALL et al., "Cerebral metabolic and cognitive decline in persons at genetic risk for Alzheimer's disease," <u>PNAS</u> , 97(11):6037-6042 (2000).			<input type="checkbox"/>
	156	SOLOMON et al., "Monoclonal antibodies inhibit in vitro fibrillar aggregation of the Alzheimer β-amyloid peptide," <u>PNAS</u> , 93:452-455 (1996).			<input type="checkbox"/>
	157	SOLOMON, B., "Immunological approaches as therapy for Alzheimer's disease," <u>Expert Opin. Biol. Ther.</u> , 2(8):907-917 (2002).			<input type="checkbox"/>
	159	STEIN et al., "Lack of Neurodegeneration in Transgenic Mice Overexpressing Mutant Amyloid Precursor Protein is Associated with Increased Levels of Transthyretin and Activation of Cell Survival Pathways," <u>The Journal of Neuroscience</u> , 22(17):7380-7388 (2002).			<input type="checkbox"/>
	160	SU et al., "Intravascular infusions of soluble β-amyloid compromise the blood-brain barrier, activate CNS Glial cells and induce peripheral hemorrhage," <u>Brain Research</u> , 818:105-107 (1999).			<input type="checkbox"/>
	161	TAKAHASHI, M. "Phosphorylation of α-synuclein characteristic of synucleinopathy lesions is recapitulated in α-synuclein transgenic Drosophila," <u>Neuroscience Letters</u> , 336: 155-158 (2003).			<input type="checkbox"/>

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Substitute for form 1449B/PTO				<b>Complete if Known</b>	
				<i>Application Number</i>	10/698,099
				<i>Filing Date</i>	October 31, 2003
				<i>First Named Inventor</i>	Schenk, Dale B.
				<i>Art Unit</i>	1648
				<i>Examiner Name</i>	Michelle S. Horning
Sheet	9	of	10	<i>Attorney Docket Number</i>	015270-008930US

NON PATENT LITERATURE DOCUMENTS					
Examiner Initials *	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.			T <sup>2</sup>
/MH/	180	TAKEDA et al., "Abnormal Distribution of the Non- $\beta$ Component of Alzheimer's Disease Amyloid Precursor/alpha-synuclein in Lewy Body Disease as Revealed by Proteinase K and Formic Acid Pretreatment," <u>Laboratory Investigation</u> , 79(9):1169-1177 (1998).			<input type="checkbox"/>
	162	TAKEDA, A. et al., "C-terminal alpha-synuclein immunoreactivity in structures other than Lewy bodies in neurodegenerative disorders," <u>Acta Neuropathol</u> , 99:296-304 (2000).			<input type="checkbox"/>
	163	TAL et al., "Complete Freund's Adjuvant Immunization Prolongs Survival in Experimental Prion Disease in Mice," <u>Journal of Neuroscience Research</u> , 71:286-290 (2003).			<input type="checkbox"/>
	164	TANAKA et al., "NC-1900, an active fragment analog of arginine vasopressin, improves learning and memory deficits induced by beta-amyloid protein in rats," <u>European J. Pharmacology</u> , 352:135-142 (1998).			<input type="checkbox"/>
	165	TENNENT et al., "Serum amyloid P component prevents proteolysis of the amyloid fibrils of Alzheimer's disease and systemic amyloidosis," <u>PNAS</u> , 92:4299-4303 (1995).			<input type="checkbox"/>
	166	TSIM, K.W. et al., "Monoclonal antibodies specific for the different subunits of asymmetric acetylcholinesterase from chick muscle," <u>J. Neurochem.</u> , 51(1):95-104 (1988).			<input type="checkbox"/>
	207	UBOL et al., "Roles of Immunoglobulin Valency and the Heavy-Chain Constant Domain in Antibody-Mediated Downregulation of Sindbis Virus Replication in Persistently Infected Neurons," <u>J Virol</u> , 1995 March; 69(3): 1990-1993.			<input type="checkbox"/>
	172	WALKER et al., "Labeling of Cerebral Amyloid <i>In Vivo</i> with a Monoclonal Antibody," <u>J. Neuropath. Exp. Neurology</u> , 53(4):377-383 (1994).			<input type="checkbox"/>
	173	WATSON et al., "Chapter 14: The Introduction of Foreign Genes into Mice," <u>Molecular Biology of Watson Recombinant DNAs</u> , 2nd ed., 255-272 (1993).			<input type="checkbox"/>
	174	WEINREB et al., "NACP, A Protein Implicated in Alzheimer's Disease and Learning, Is Natively Unfolded," <u>Biochemistry</u> , 35(43):13709-13715 (1996).			<input type="checkbox"/>



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<b>NON PATENT LITERATURE DOCUMENTS</b>			
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/MH/	175	WISNIEWSKI et al., "Therapeutics in Alzheimer's and Prion Diseases," <u>Biochemical Society Transactions</u> , 30(4):574-587 (2002).	<input type="checkbox"/>
/MH/	210	WONG et al., "Neuritic Plaques and Cerebrovascular Amyloid in Alzheimer Disease are Antigenically Related," <u>PNAS</u> , 82:8729-8732 (1985).	<input type="checkbox"/>
/MH/	176	YOSHIMOTO et al., "NACP, the precursor protein of the non-amyloid $\beta$ /A4 protein (A $\beta$ ) component of Alzheimer disease amyloid, binds A $\beta$ and stimulates A $\beta$ aggregation," <u>PNAS</u> , 92:9141-9145 (1995).	<input type="checkbox"/>
/MH/	211	ZHOU et al., "A Human Single-Chain Fv Intrabody Blocks Aberrant Cellular Effects of Overexpressed alpha-Synuclein," <u>Mol Ther.</u> , 10(6):1023-31 (2004).	<input type="checkbox"/>

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